

net Explorer					7 Divir - 2 Sequence (203~390) 1 - 282 : Signal peptide + Propeptide	→ SEQ ID No:3 of the present invention: 2-18 of the above BMP-2 (red underline): 17 amino acids	→ SEQ ID No:6 of the present invention: 24-40 of the above BMP-2(pink	underline) : 17 amino acids		•SEQ ID No: 10 of the present invention: 283-302 of BMP- 2 precusor (violet underline)	• SEQ ID No: 11 of the present invention: 355-374 of BMP-2 precusor (green underline)		 SEQ ID No: 12 of the present invention: 370-390 of BMP-2 precusor (yellow underline)
derosoft Inter				Tools	Blast		6 <u>0</u> SEFELRLLSM	$12\frac{0}{0}$ SFHHEESLEE	18 <u>0</u> INIYEIIKPA	24 <u>0</u> ahleekogvs	30 <u>0</u> RLKSSCKRHP	36 <u>0</u> SVNSKIPKAC	
(Aluman) — 1				n Mass (Da)	3 44,702		5 <u>0</u> PSSQPSDEVL	11 <u>0</u> RAASRANTVR	17 <u>0</u> LGNNSSFHHR	23 <u>0</u> HANHGFUVEV	29 <u>0</u> KROAKHKORK	35 <u>0</u> NHAIVQTLVN	
– Homo sapiens 도움말(!!)	Q)			Length	FASTA 396		4 <u>0</u> RKFAAASSGR	10 <u>0</u> GSPAPDHRLE	16 <u>0</u> OVFREQMODA	22 <u>0</u> Pavmrutaqg	28 <u>0</u> GKGHPLHKRE	34 <u>0</u> FPLADHLNST	EGGGCR
90gursor - 那 5子(I) - 도움	검색 🦙 즐겨찾기	12643			∄	/ersion 1.	3 <u>0</u> Aaglvpelgr	9 <u>0</u> DLYRRHSGQP	15 <u>0</u> Teefitsael	21 <u>0</u> ASRWESFDVT	27 <u>0</u> RPLLVTFGHD	33 <u>0</u> HAFYCHGECP	39 <u>0</u> VLKNYODWV
protein 2 pr 즐거찾기(A)		ot.org/uniprot/F			iParc].	tober 1, 1989. \ 3A3987B25E60	2 <u>0</u> LLLPQVLLGG	8 <u>0</u> RDAVVPPYML	14 <u>0</u> RFFFNLSSIP	20 <u>0</u> LLDTRLVNON	26 <u>0</u> HODEHSWSOI	32 <u>0</u> NDWIVAPPGY	38 <u>0</u> Mlyldenekv
경) Bone morphogenetic protein 2 precursor - Homo sapiens (Human) - Microsoft Internet Explorer 파일氏) 편집(E) 보기(Y) 즐거찾기(A) 도구(I) 도움말(H)		(존] http://www.uniprot.org/uniprot/P12643	To the state of th	Sequence	P12643-1 [UniParc]	Last modified October 1, 1989. Version 1. Checksum: 20653A3987B25E60	1 <u>0</u> MVAGTRCLLA	7 <u>0</u> FGLKQRPTPS	13 <u>0</u> LPETSGKTTR	19 <u>0</u> Tanskfpvtr	25 <u>0</u> KRHVRISRSL	31 <u>0</u> LYVDFSDVGU	37 <u>0</u> CVPTELSAIS
2] Bone m IPS(E) 语	◎ 뒤로 •	74(D)											

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osoit internet Explorer	\$\$ \$\$			Graphical view						Mary Mary	John Mrs.		AP OF	100 200	APP COMP	man data	
- Homo sapiens (Human) – Microsoft Internet Explorer 도움말(出)				Description		Potential		Bone morphogenetic protein 2		N-linked (GlcNAc) [Potential]	N-linked (GlcNAc) Potential	N-linked (GlcNAc) Potential	N-linked (GICNAc) Potential				Interchain
protein 2 precursor - 즐겨찾기(ል) 도구(I)		/uniprot/P12643	or (Termins)	Position(s) Length		1-23 23	24 - 282 259	283 – 396 114		135	163	164	200 1	296 ↔ 361	325 ↔ 393	329 ↔ 395	360
🤔 Bone morphogenetic protein 2 precursor - Ho 파일(E) 편집(E) 보기(Y) 즐겨찾기(A) 도구(I) 도움말	지 (平全(D) (圖 http://www.uniprot.org/uniprot/P12643	West and a second of (T)	Feature key Pos	Molecule processing	☐ Signal peptide	☐ Propeptide 2	☐ Chain 28	Amino acid modifications	☐ Glycosylation	☐ Glycosylation	Glycosylation	☐ Glycosylation	Disulfide bond 296	☐ Disulfide bond 32₹	Disulfide bond	Disulfide bond



→ SEQ ID No: 4 of the present invention: 2-18 of the above BMP-4 (red underline): 17 • SEQ ID No: 15 of the present invention: SEQ ID No: 16 of the present invention: •SEQ ID No: 14 of the present invention: →1~292 : Signal peptide + Propeptide 293-313 of BMP- 4 precusor (violet 366-386 of BMP-4 precusor (green →BMP-4 sequence (293~408). Ņ, e morphogenetic protein 4 precursor – Homo sapiens (Human) – Microsoft Internet Explorer amino acids underline) underline) 1<u>0</u> 2<u>0</u> 3<u>0</u> 4<u>0</u> 5<u>0</u> 6<u>0</u> MIPGNRNLMV VLLCOVLLGG ASHASLIPET GKKKVAEIQG HAGGRRSGQS HELLRDFEAT 190 200 210 220 220 240 YEVMKPPAEV VPGHLITRLI DIRLVHHNVI RWETFDVSPA VLRWTREKOP NYGLAIEVTH 360 $\frac{70}{\text{LLQMFGLRRR}} \frac{80}{\text{POPSKSAVIP}} \frac{90}{\text{DYMRDLYRLQ}} \frac{90}{\text{SGEEEEEQIH}} \frac{110}{\text{STGLEYPERP}} \frac{120}{\text{ASRANTVRSF}}$ 13<u>0</u> 14<u>0</u> 15<u>0</u> 16<u>0</u> 17<u>0</u> 18<u>0</u> HHEEHLENIP GTSENSAFRF LFNLSSIPEN EVISSAELRL FREQVDQGPD WERGFHRINI 31<u>0</u> 32<u>0</u> 33<u>0</u> 34<u>0</u> 35<u>0</u> 36<u>0</u> APKKENIZMERR, HSLAU</u>DFSDV GUNDUIVAPP GYQAFYCHGD CPFPLADHLN STNHAIVQTL Diast 25<u>0</u> 26<u>0</u> 27<u>0</u> 28<u>0</u> 29<u>0</u> LHOTRTHQGQ HVRISRSLPQ GSGNWAQLRP LLVTFGHDGR GHALTRRRRA KR KVVLKNYQEM VVEGCGCR QD Ż 400 (1) として ○ ★ ○ 日8時(日) 390 CHIEFT DE YD Last modified October 1, 1989. Version 1. 보기(火) 즐거찾기(A) 도구(T)) (http://www.uniprot.org/uniprot/P12644 Checksum: 79B01179DBB98204 380 12044-110mm arch 370 THE WASHING * Hide 园区(0) <u>щ</u>

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382-402 of BMP-4 precusor (yellow

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sor – Homo sapiens (Human) – Microsofi Internet Explorer (I) 도움말(H)	次 ē 3 4 ≠ 7 € 6 6 · □ 6 8 8			Length Description		19 Potential	273	116 Bone morphogenetic protein 4		1 N-linked (GlcNAc) Potential	1 N-linked (GlcNAc) Potential	1 N-linked (GlcNAc) Potential	1 N-linked (GlcNAc) Potential	By similarity	By similarity	By similarity	Interchain By similarity
protein 4 preceurso 즐겨찾기(8) 도구(T)	(*) (*) 전 (*) 즐거찾기	ıt,org/uniprot/P12644		Position(s) Le		1-19	20 – 292	293 – 408	SI	143	208	350	365	308 ↔ 373	337 ↔ 405	341 ↔ 407	372
$^{f 2}$ Bone morphogenetic protein 4 precursor — Homo m일($f E$) 편집($f E$) 보기($f V$) 즐거찾기($f A$) 도구($f I$) 도움말($f H$)	<u>*</u>	주소(D) (월) http://www.uniprot.org/uniprot/P12644	Continue of the state of the st	Feature key	Molecule processing	☐ Signal peptide	☐ Propeptide	□ Chain	Amino acid modifications	☐ Glycosylation	Glycosylation	Glycosylation	Glycosylation	☐ Disulfide bond	Disulfide bond	Disulfide bond	Disulfide bond



2-18 of the above BMP-6 (red underline): SEQ ID No: 19 of the present invention: SEQ ID No: 20 of the present invention: → SEQ ID No: 5 of the present invention: •SEQ ID No: 18 of the present invention: → 1~374: Signal peptide + Propeptide 487-510 of BMP-6 precusor (yellow 472-490 of BMP-6 precusor (green 397-418 of BMP- 6 precusor (violet →BMP-6 sequence (375~513) Ŋ 17 amino acids 🗿 Bone morphogenetic protein 6 precursor - Homo sapiens (Human) - Microsoft Internet Explorer underline) underline) underline) 8 > 42<u>0</u> ACRKHELYVS 7<u>0</u> 8<u>0</u> 9<u>0</u> 10<u>0</u> 11<u>0</u> 12<u>0</u> OSSSGFLYRR LKTCEKRENQ KEILSVLGLP HRPRPLHGLQ OPQPPALRQQ EEQQQQQQLP 130 140 150 160 160 170 180 180 RGEPPFGRLK SAPLFMLDLY NALSADNDED GASEGERQOS UPHEAASSSQ RRQPPPGAAH 19<u>2</u> 20<u>0</u> 21<u>0</u> 22<u>0</u> 23<u>0</u> 24<u>0</u> PLNRKSLLAP GSGSGASPL TSAQDSAFLN DADNVMSFVN LVEYDKEFSP RORHHKEFKF 25<u>0</u> 25<u>0</u> 25<u>0</u> 30<u>0</u> 30<u>0</u> 30<u>0</u> NLSQIPEGEV VTAAEFRIYK DCVMGSFKNQ TFLISIYQVL QEHQHRDSDL FLLDTRVVWA 10 $2\underline{0}$ $3\underline{0}$ $3\underline{0}$ $6\underline{0}$ MPGLGRRAQU LCWUWGLLCS CCGPPPLRPP LPAAAAAAG GQLLGDGGSP GRTEQPPPSP 310 320 350 340 350 350 350 SEEGULEFDI TATSNLUVVT POHNMGLQLS VVTRDGVHVH PRAAGLVGRD GPYDKQPFMV 43<u>0</u> 44<u>0</u> 45<u>0</u> 45<u>0</u> 46<u>0</u> 46<u>0</u> 47<u>0</u> 480 FQDLGWQDWI IAPKGYAANY CDGECSFPLN AHMNATNHAI VQTLVHLMNP KYYPKPGCAP Blast Tools 4. 01. 01. SDYNSSELKT Mass (Da) 57,226 Length 513 400 SODVARVSSA (i) FASTA H C C H 12% 雪牙茶7 五岛四(田) 37<u>0</u> 38<u>0</u> 39<u>0</u> AFFKVSEVHV RTTREASSRR RQOSRNRSTQ 50<u>0</u> 51<u>0</u> FDDNSNVILK KYRNMYVRAC Last modified August 1, 1991. Version 1. 平全(D) (图) http://www.uniprot.org/uniprot/P22004 Checksum: 3F19155B36049278 @'∀찾기(₫) ্ট ডি P22004-1 [UniParc] 49<u>0</u> TKLNAISVLY 以 以 以 ***** Sequence 면접(E) ٠ 印品自

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- Homo sapiens (Human) - Microsoft Internet Explorer 도움말(出)				Description		Potential	Potential	Bone morphogenetic protein 6		N-linked (GlcNAc) Potential	N-linked (GlcNAc) Potential	N-linked (GloNAc) Potential	N-linked (GlcNAc) [Potential]	N-linked (GlcNAc) [Potential]	By simulanty	By similarity	Bysimilarity	Interchain By similarity		R → C; dbSNP rs10458105.
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groteln Signese 즐거찾기(A) 도		ot, org/uniprot/P22(1-20	21 – 374	375-513	JS.	241	269	386	404	454	412 ↔ 478	441 ↔ 510	445 ↔ 512	477	٠	257
클 Bone morphogenetic protein 6 precursor — Homo s 파일(5) 편집(5) 보기(y) 즐거찾기(A) 도구(I) 도움말(H)] × ○ • ≥H €	주소(D) (國 http://www.uniprot,org/uniprot/P22004	Sequence annoration (Features)	Feature key	Molecule processing	☐ Signal peptide	☐ Propeptide	□ Chain	Amino acid modifications	☐ Glycosylation	Glycosylation	Glycosylation	☐ Glycosylation	Glycosylation	Disulfide bond	Disulfide bond	☐ Disulfide bond	Disulfide bond	Natural variations	Natural variant



🏂 Bone morphogenetic protein 7 precursor – Homo sapiens (Human) – Microsoft Internet Explorer

도움말(比) (<u>1</u>) 즐거찾기(A) 五 万 () 면접(E) 田智(日)

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431 FASTA

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Last modified November 1, 1990. Version 1. Checksum: 47AD5E45C6815F8A 10 20 30 4<u>0</u> 5<u>0</u> 6<u>0</u> mhvrslraaa phsfvalwap lflirsalad fsldnevhss fihrrlrsqe rremoreils

25<u>0</u> 26<u>0</u> 27<u>0</u> 28<u>0</u> 29<u>0</u> 30<u>0</u> HNLGLQLSVE TLDGQSINPK LAGLIGRHGP ONKQPFMVAF FKATEVHFRS INSTGSKQRS

ONRSKIPKNO EALRMANVAE NSSSDOROAC KKHELYVSFR DLGUQDUIIA PEGYAAYYCE 340

37<u>0</u> 38<u>0</u> 39<u>0</u> 40<u>0</u> 41<u>0</u> 42<u>0</u> GECAFPLNSY MNATNHAIVQ TLVHFINPE<u>T VPKPCCAPTQ LNAISVLYF</u>D DSSNVILKKY

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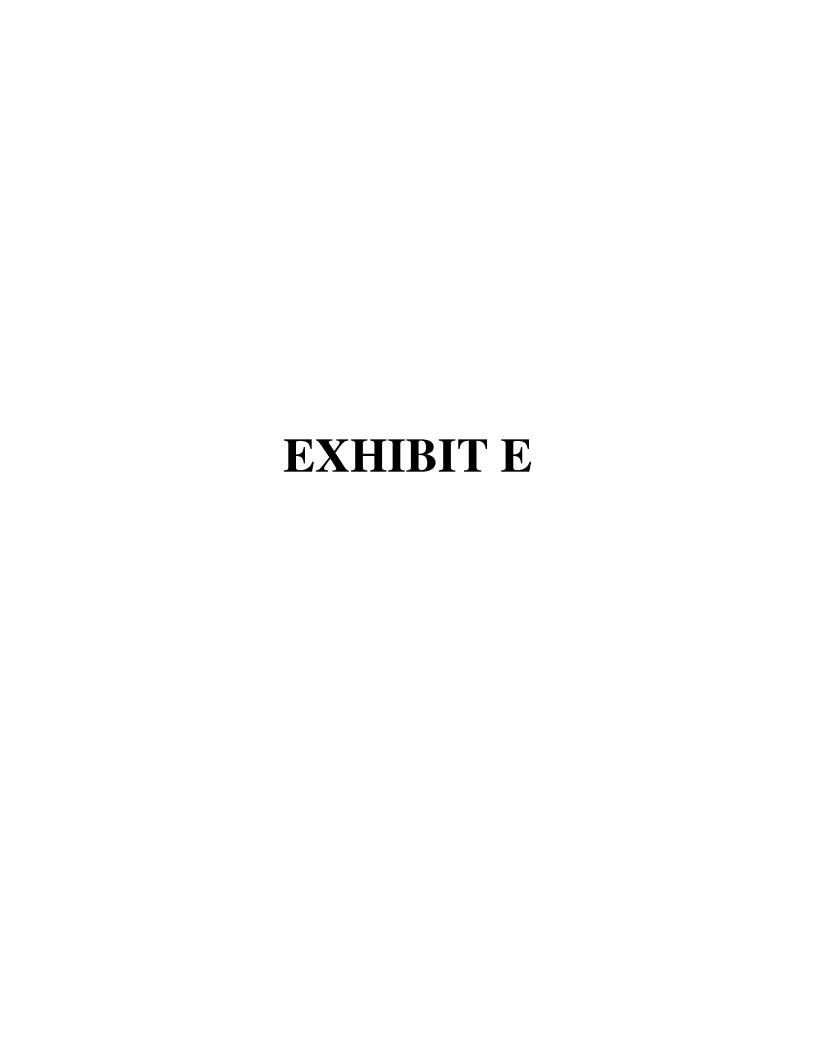
RNMVVRACGC

→ 1~292: Signal peptide + Propeptide →BMP-7 sequence (293~431)

No: 23 of the present invention (390-409 of → SEQ ID No: 21 of the present invention: (The above sequence is equal to SEQ ID 98-117 of BMP-7 (pink underline) BMP-7 precusor.)

*SEQ ID No: 22 of the present invention: 320-340 of BMP-7 precusor (violet underline)

405-423 of BMP-7 precusor (red underline) SEQ ID No: 24 of the present invention:



Partial translation for KR 10-2004-0019010

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[Disclosure of invention]

<15> To achieve the above object, the present invention provides a barrier membrane and a implant which have a cell adhesion-inducing peptide and/or tissue growth factor-derived peptide immobilized on the surface of the membrane or the implant bonded with a cross-linking agent.

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<19> Specifically, the cell adhesion-inducing peptide is preferably a peptide having an amino acid sequence of SED ID NO: 1. More preferably, it is an amino acid sequence of SEQ ID NO: 2 or an amino acid sequence of SEQ ID NO: 3 designed to maintain the structural stability of the amino acid sequence of SEQ ID NO: 1, RGD. Furthermore, the tissue growth factor-derived peptide is a peptide identified and chemically synthesized from the active site of the tissue growth factor. Preferably it is at least one peptide selected from the group consisting of the following peptides:

<20> (a) the amino acid sequence at positions 283-302 (SEQ ID NO: 4), the amino acid sequence at positions 335-353 (SEQ ID NO: 5) and the amino acid sequence at positions 370-390 (SEQ ID NO: 6) of bone morphogenetic proteins (BMP)-2;

<21> the amino acid sequence at positions 293-313 (SEQ ID NO: 7), the amino acid sequence at positions 360-379 (SEQ ID NO: 8) and the amino acid sequence at positions 382-402 (SEQ ID NO: 9) of BMP-4;

<22> the amino acid sequence at positions 397-418 (SEQ ID NO: 10), the amino

acid sequence at positions 472-490 (SEQ ID NO: 11) and the amino acid sequence at positions 487-510 (SEQ ID NO: 12) of BMP-6;

<23> the amino acid sequence at positions 320-340 (SEQ ID NO: 13), the amino acid sequence at positions 390-409 (SEQ ID NO: 14) and the amino acid sequence at positions 405-423 (SEQ ID NO: 15) of BMP-7;

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<24> (b) the amino acid sequence at positions 199-204 (SEQ ID NO: 16), the amino acid sequence at positions 151-158 (SEQ ID NO: 17), the amino acid sequence at positions 275-291 (SEQ ID NO: 18), the amino acid sequence at positions 20-28 (SEQ ID NO: 19), the amino acid sequence at positions 65-90 (SEQ ID NO: 20), the amino acid sequence at positions 150-170 (SEQ ID NO: 21) and the amino acid sequence at positions 280-290 (SEQ ID NO: 22) of bone sialoprotein,

<25> (c) the amino acid sequence at positions 242-250 (SEQ ID NO: 23), the amino acid sequence at positions 279-299 (SEQ ID NO: 24) and the amino acid sequence at positions 343-361 (SEQ ID NO: 25) of a transforming growth factor beta 1,

<26> (d) the amino acid sequence at positions 100-120 (SEQ ID NO: 26) and the amino acid sequence at positions 121-140 (SEQ ID NO: 27) of a platelet-derived growth factor,

<27> (e) the amino acid sequence at positions 23-31 (SEQ ID NO: 28) and the amino acid sequence at positions 97-105 (SEQ ID NO: 29) of an acidic fibroblast growth factor,

<28> (f) the amino acid sequence at positions 16-27 (SEQ ID NO: 30), the amino acid sequence at positions 37-42 (SEQ ID NO: 31), the amino acid sequence at positions

78-84 (SEQ ID NO: 32) and the amino acid sequence at positions 107-112 (SEQ ID NO: 33) of a basic fibroblast growth factor,

<29> (g) the amino acid sequence at positions 255-275 (SEQ ID NO: 34), the amino acid sequence at positions 475-494 (SEQ ID NO: 35) and the amino acid sequence at positions 551-573 (SEQ ID NO: 36) of dentin sialoprotein,

<30> (h) the amino acid sequence at positions 63-83 (SEQ ID NO: 37), the amino acid sequence at positions 84-103 (SEQ ID NO: 38), the amino acid sequence at positions 104-116 (SEQ ID NO: 39) and the amino acid sequence at positions 121-140 (SEQ ID NO: 40) of a heparin binding EGF-like growth factor,

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<31> (i) the amino acid sequence at positions 326-350 (SEQ ID NO: 41), the amino acid sequence at positions 351-371 (SEQ ID NO: 42), the amino acid sequence at positions 372-400 (SEQ ID NO: 43), the amino acid sequence at positions 401-423 (SEQ ID NO: 44), the amino acid sequence at positions 434-545 (SEQ ID NO: 45), the amino acid sequence at positions 546-651 (SEQ ID NO: 46), the amino acid sequence at positions 1375-1433 (SEQ ID NO: 47), the amino acid sequence at positions 1435-1471 (SEQ ID NO: 48), the amino acid sequence at positions 1475-1514 (SEQ ID NO: 49), the amino acid sequence at positions 1515-1719 (SEQ ID NO: 50), the amino acid sequence at positions 1764-1944 (SEQ ID NO: 51) and the amino acid sequence at positions 2096-2529 (SEQ ID NO: 52) of the cadherin EGF LAG seven-pass G-type receptor 3,

<32> (j) the amino acid sequence at positions 54-159 (SEQ ID NO: 53), the amino acid sequence at positions 160-268 (SEQ ID NO: 54), the amino acid sequence at positions 269-383 (SEQ ID NO: 55), the amino acid sequence at positions 384-486

(SEQ ID NO: 56) and the amino acid sequence at positions 487-612 (SEQ ID NO: 57) of an osteoblast specific cadherin (OB-cadherin).

<33> The N-terminal end of the peptide has an addition of cysteine and two glycines so as to stabilize the structure of the peptide and to facilitate chemical immobilization of the peptide to the barrier membrane.

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<35> As the barrier membrane to be surface-activated by the present invention, all kinds and types of barrier membranes can be used if they are used in the technical field. Pre Preferred examples of these barrier membranes include porous membranes made of polylactic acid, regeneraton membranes made of nanofibers of chitin or chitosan, and film-shaped barrier membranes made of chitin or chitosan. Also, as the implants, titanium implants are preferably used but are not limited thereto. In this respect, the surface of the implants is preferably modified by oxidation and nitrification so as to facilitate the adhesion of the active peptide to the surface.

VERIFICATION OF TRANSLATION

I, BAE, Young Sim of 11th Yeosam Bldg., 648-23, Yeoksam-dong, Gangnam-gu, Seoul, 135-080, Republic of Korea

State that the attached document is a true and accurate translation of Korean Patent Application No. 10-2004-0019010 in Korean_(language of original specification) to the best of my knowledge and belief.

Dated this August 3, 2009

Signature of Translator: